

(12) INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

(19) World Intellectual Property Organization
International Bureau(43) International Publication Date
23 October 2003 (23.10.2003)

PCT

(10) International Publication Number
WO 03/086382 A1(51) International Patent Classification⁷: **A61K 31/336**,
38/04, 9/00

(21) International Application Number: PCT/US03/10976

(22) International Filing Date: 10 April 2003 (10.04.2003)

(25) Filing Language: English

(26) Publication Language: English

(30) Priority Data:
60/371,791 11 April 2002 (11.04.2002) US
60/414,705 30 September 2002 (30.09.2002) US(71) Applicant (for all designated States except US): **CHIL-
DREN'S MEDICAL CENTER CORPORATION**
[US/US]; 300 Longwood Avenue, Boston, MA 02115
(US).

(72) Inventors; and

(75) Inventors/Applicants (for US only): **SATCHI-
FAINARO, Ronit** [IL/US]; 129 Sherman Road, Chestnut
Hill, MA 02467 (US). **FOLKMAN, Judah** [US/US]; 18
Chatham Circle, Brookline, MA 02115 (US).(74) Agents: **RESNICK, David, S.** et al.; Nixon Peabody LLP,
101 Federal Street, Boston, MA 02110 (US).(81) Designated States (national): AE, AG, AL, AM, AT, AU,
AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU,
CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC,
LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW,
MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SC, SD, SE,
SG, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ,
VC, VN, YU, ZA, ZM, ZW.(84) Designated States (regional): ARIPO patent (GH, GM,
KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW),
Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM),
European patent (AT, BE, BG, CH, CY, CZ, DE, DK, EE,
ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO,
SE, SI, SK, TR), OAPI patent (BF, BJ, CF, CG, CI, CM,
GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG).**Published:**

- with international search report
- before the expiration of the time limit for amending the claims and to be republished in the event of receipt of amendments

For two-letter codes and other abbreviations, refer to the "Guidance Notes on Codes and Abbreviations" appearing at the beginning of each regular issue of the PCT Gazette.

(54) Title: TNP-470 POLYMER CONJUGATES AND USE THEREOF

(57) **Abstract:** The present invention relates to conjugates of water-soluble polymers and o(chloroacetyl-carbamoyl) fumagillol (TNP-470) and use of those conjugates as specific intracellular carriers of the TNP-470 into tumor vessels. The present invention further relates to use of those conjugates to lower the neurotoxicity of TNP-470. Preferably, the polymer has a molecular weight in the range of 100 Da to 800 kDa. More preferably, the polymer has a molecular weight no greater than 60 kDa. Most preferably, the polymer has a molecular weight in the range of 15 kDa to 40 kDa.

WO 03/086382 A1